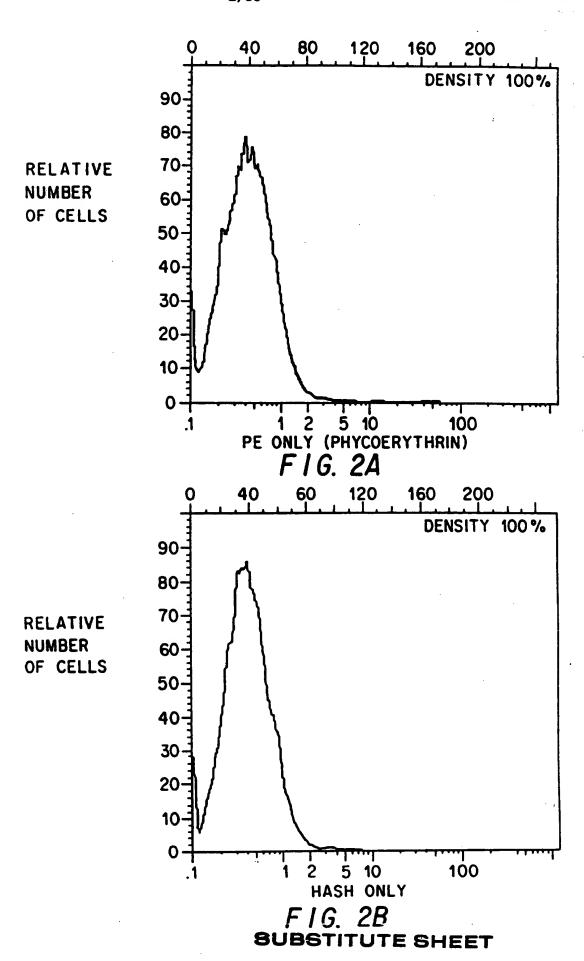
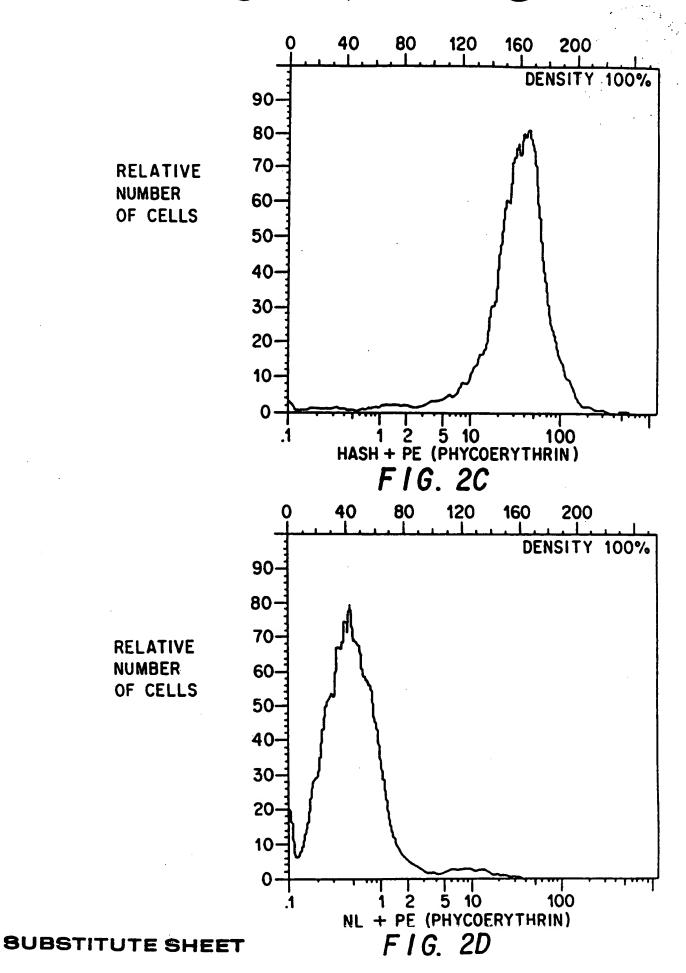
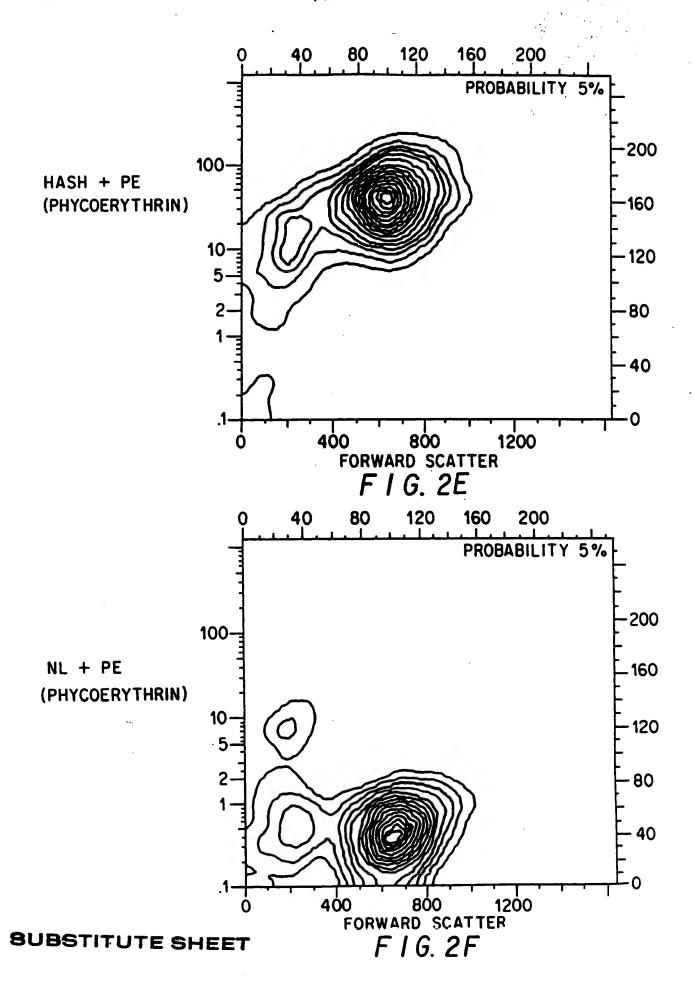
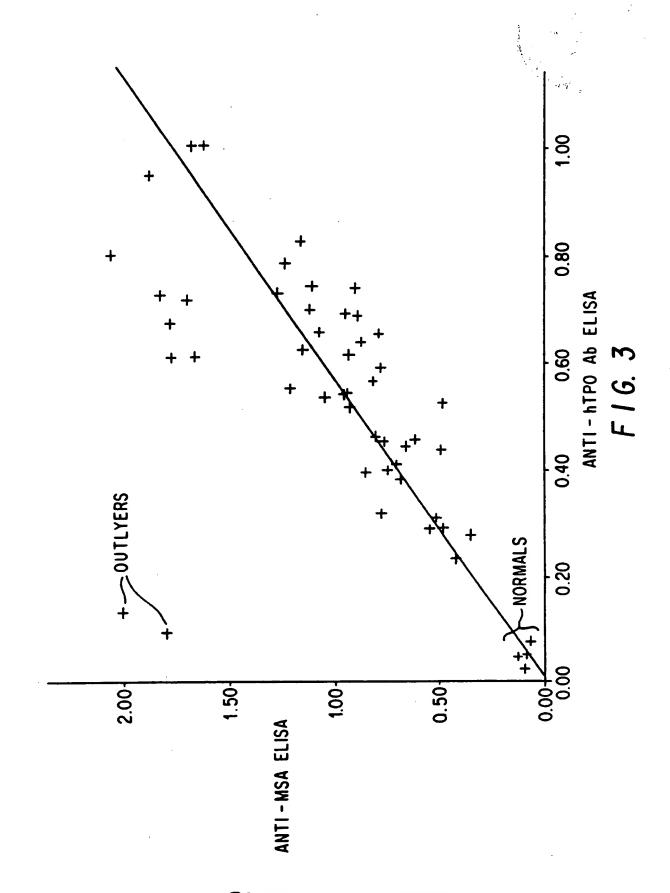


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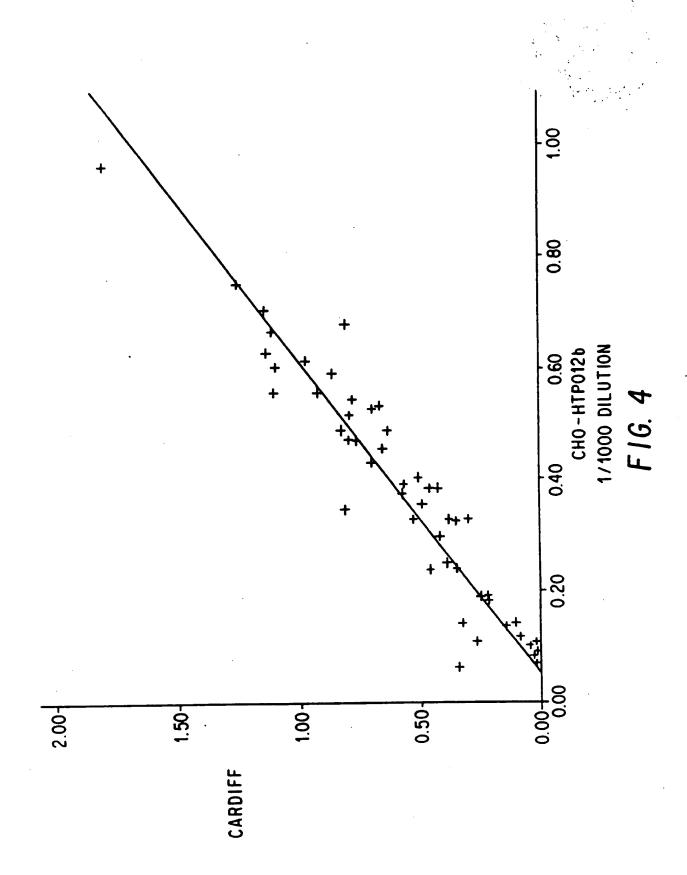




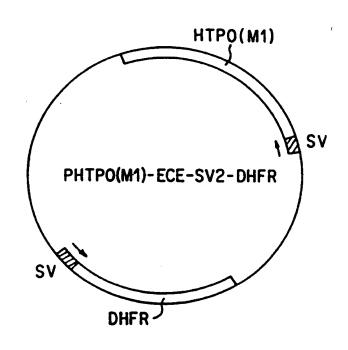




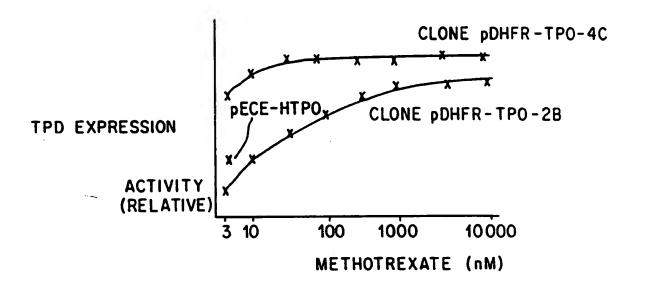
**SUBSTITUTE SHEET** 



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F1G. 8



F1G. 5

2611	5992	€.
AGG CTC CCT CGG GTG ACT TGG ATC IC	GTG ACT TGG ATC ICC ATG TCG CTG GCT GCT CTG CTG ATC G	pHTPO-BS
Eco RI		
Stop	Stop	
AGG CTC CCT CGG GTG ACT TGA AT <u>I C</u> C	GTG ACT TGA AT <u>i c</u> cc atg t <u>a</u> g ctg gct gct ctg ctg atc g	pHTPD(M1)-BS
	F 1 G. 6	

GAG	ĢСА	ATT	GAG	GCG	CCC	ATT	TCA	27 <b>G</b> AA	GAG	TTA	CAĢ	CCG	TGA	AAA	TTA	СТС	54 AGC
AGT	GCA	GTT	GGC	TGA	GAA	GAG	GAA	81 AAA	AGA					GCT Ala		_	
GTC Val																	_
AAA Lys																	
GAA Glu																	
AAG Lys																	
GAG Glu	CCA Pro	ACA Thr	AGC Ser	GGA Gly	GTG Val	ATT lle	GCC Ala	351 CGA Arg	GCA Ala	GCA Ala	GAG Glu	ATA Ile	ATG Met	GAA Glu	ACA Thr	TCA Ser	378 ATA
CAA G l n	GCG Ala	ATG Met	AAA Lys	AGA Arg	AAA Lys	GTC Val	AAC Asn	405 CTG Leu	AAA Lys	ACT Thr	CAA Gln	CAA Gln	TCA Ser	CAG Gln	CAT His	CCA Pro	432 ACG Thr
GAT *	GCT Ala	TTA Leu	TCA Ser	GAA Glu	GAT Asp	CTG Leu	CTG Leu	459 AGC Ser	ATC Ile	ATT Ile	GCA Ala	AAC Asn	ATG Met	TCT Ser	GGA Gly	TGT Cys	486 CTC Leu

# FIG. 7A

513 540 CCT TAC ATG CTG CCC CCA AAA TGC CCA AAC ACT TGC CTG GCG AAC AAA TAC AGG Pro Tyr Met Leu Pro Pro Lys Cys Pro Asn Thr Cys Leu Ala Asn Lys Tyr Arg 594 CCC ATC ACA GGA GCT TGC AAC AAC AGA GAC CAC CCC AGA TGG GGC GCC TCC AAC Pro Ile Thr Gly Ala Cys Asn Asn Arg Asp His Pro Arg Trp Gly Ala Ser Asn 621 648 ACG GCC CTG GCA CGA TGG CTC CCT CCA GTC TAT GAG GAC GGC TTC AGT CAG CCC Thr Ala Leu Ala Arg Trp Leu Pro Pro Val Tyr Glu Asp Gly Phe Ser Gln Pro 702 675 CGA GGC TGG AAC CCC GGC TTC TTG TAC AAC GGG TTC CCA CTG CCC CCG GTC CGG Arg Gly Trp Asn Pro Gly Phe Leu Tyr Asn Gly Phe Pro Leu Pro Pro Val Arg 729 756 GAG GTG ACA AGA CAT GTC ATT CAA GTT TCA AAT GAG GTT GTC ACA GAT GAT GAC Glu Val Thr Arg His Val Ile Gln Val Ser Asn Glu Val Val Thr Asp Asp Asp 810 783 CGC TAT TCT GAC CTC CTG ATG GCA TGG GGA CAA TAC ATC GAC CAC GAC ATC GCG Arg Tyr Ser Asp Leu Leu MET Ala Trp Gly Gln Tyr Ile Asp His Asp Ile Ala 864 837 TTC ACA CCA CAG AGC ACC AGC AAA GCT GCC TTC GGG GGA GGG TCT GAC TGC CAG Phe Thr Pro Gln Ser Thr Ser Lys Ala Ala Phe Gly Gly Ser Asp Cys Gln 918 891 ATG ACT TGT GAG AAC CAA AAC CCA TGT TTT CCC ATA CAA CTC CCG GAG GAG GCC Met Thr Cys Glu Asn Gln Asn Pro Cys Phe Pro Ile Gln Leu Pro Glu Glu Ala 972 945 CGG CCG GCC GCG GGC ACC GCC TGT CTG CCC TTC TAC CGC TCT TCG GCC GCC TGC Arg Pro Ala Ala Gly Thr Ala Cys Leu Pro Phe Tyr Arg Ser Ser Ala Ala Cys

## FIG. 7B

								999								1	026
GGC	ACC	GGG	GAC	CAA	GGC	GCG	CTC		ดดด	AAC	CTG	TCC	ACG	GCC	AAC		
			Asp														
•		•	•		•				_	×							
								053	CTC	C 4 C		TOO	400	CTC	<b>TAT</b>		080
CAG	CAG	ATG	AAC	GGG	116	ACC	ICu	HU	LIU	GAL	ulu	ILL	ALL	<b>UIU</b>	IAL	սսե Մ	AGC
ն (n	<u>ն</u> Լո	Met	Asn	ціу	Leu	ınr	26L	rne	Leu	ASP	Ala	3er	ınr	ναι	ıyr	υιу	76L
							1	107								1	134
TCC	ררה	הרר	CTA	האה	ΔGG	CAG			AAC	TGG	ACC	AGT	GCC	GAA	666	-	
Ser.	Pro	Aln	Leu	Glu	Ara	Gln	Leu	Ara	Asn	Trp	Thr	Ser	Ala	Glu	Gly	Leu	Leu
JC.				• • • •	'- Э	•	-	3	×	F					,		
							1	161								1	188
CGC	GTC	CAC	GGC	CGC	CTC	CGG			GGC	CGC	GCC	TAC	CTG	CCC	TTC		
Ara	Val	HIS	Gly	Arg	Leu	Arg	Asp	Ser	Gly	Arg	Ala	Tyr	Leu	Pro	Phe	Val	Pro
J			•						-	_							
								1215	C.A.E.	000		446	ccc	CC 4	C 4 C		1242
CCA	CCC	GCG	CCT	GCG	GCC	161	GCU	רבי	6A6	Dun	Lili	AAL	Pag	LUA	UAU C.L.	ALL	\u\ \
Pro	Arg	Ala	Pro	Ala	Ala	Lys	Ala	Pro	ษเน	Pro	uty	ASN	Fro	uty	utu	I FIF	Hry
							,	1269									1296
ההה	ררר	דהר	TTC	CTG	GCC	GGA			CGC	GCC	AGC	GAG	GTC	CCC	TCC		
li l v	Pro	Cvs	Phe	Leu	Ala	Glv	ASD	Gly	Arg	Ala	Ser	Glu	Val	Pro	Ser	Leu	Thr
U ( )		0,2				- ,	•	•	. •								
								1323									1350
GCA	CTG	CAC	ACG	CTG	TGG	CTG	CGC	GAG	CAC	AAC	CCC	CTG	GCC	GCG	GCG	CIC	AAG
Ala	Leu	His	Thr	Leu	Trp	Leu	Arg	Glu	His	Asn	Arg	Leu	Ala	Ala	Ala	Leu	Lys
								1377									1404
CCC	CTC	AAT	GCG	LVL	TGG	۸۲۲			הרר	הדה	TAC	CAG	GAG	GCG	CGC		
410	100	AAI	Δla	Hic	Trn	Ser	Ala	Asn	Ala	Val	Tvr	Gln	Glu	Ala	Arq	Lys	Val
m (a	LEU	moli	пи	1113	р	50,		р			. ,				3	,	
								1431								_	1458
GTG	GGC	GCT	CTG	CAC	CAG	ATC	ATC	ACC	CTG	AGG	GAT	TAC	ATC	CCC	AGG	ATC	CTG
Val	Gly	Ala	Leu	His	Gln	Ile	Ile	Thr	Leu	Arg	Asp	Tyr	He	Pro	Arg	Ile	Leu

# FIG. 7C

1485 1512 GGA CCC GAG GCC TTC CAG CAG TAC GTG GGT CCC TAT GAA GGC TAT GAC TCC ACC Gly Pro Glu Ala Phe Gln Gln Tyr Val Gly Pro Tyr Glu Gly Tyr Asp Ser Thr 1539 1566 GCC AAC CCC ACT GTG TCC AAC GTG TTC TCC ACA GCC GCC TTC CGC TTC GGC CAT Ala Asn Pro Thr Val Ser Asn Val Phe Ser Thr Ala Ala Phe Arg Phe Gly His 1593 1620 GCC ACG ATC CAC CCG CTG GTG AGG AGG CTG GAC GCC AGC TTC CAG GAG CAC CCC Ala Thr Ile His Pro Leu Val Arg Arg Leu Asp Ala Ser Phe Gln Glu His Pro 1647 1674 GAC CTG CCC GGG CTG TGG CTG CAC CAG GCT TTC TTC AGC CCA TGG ACA TTA CTC ASP Leu Pro Gly Leu Trp Leu His Gln Ala Phe Phe Ser Pro Trp Thr Leu Leu 1701 1728 CGT GGA GGT GGT TTG GAC CCA CTA ATA CGA GGC CTT CTT GCA AGA CCA GCC AAA Arg Gly Gly Gly Leu Asp Pro Leu Ile Arg Gly Leu Leu Ala Arg Pro Ala Lys 1755 1782 CTG CAG GTG CAG GAT CAG CTG ATG AAC GAG GAG CTG ACG GAA AGG CTC TTT GTG Leu Gln Val Gln Asp Gin Leu Met Asn Glu Glu Leu Thr Glu Arg Leu Phe Val 1809 1836 CTG TCC AAT TCC AGC ACC TTG GAT CTG GCG TCC ATC AAC CTG CAG AGG GGC CGG Leu Ser Asn Ser Ser Thr Leu Asp Leu Ala Ser Ile Asn Leu Gin Arg Gly Arg 1863 1890 GAC CAC GGG CTG CCA GGT TAC AAT GAG TGG AGG GAG TTC TGC GGC CTG CCT CGC Asp His Gly Leu Pro Gly Tyr Asn Glu Trp Arg Glu Phe Cys Gly Leu Pro Arg 1944 1917 CTG GAG ACC CCC GCT GAC CTG AGC ACA GCC ATC GCC AGC AGG AGC GTG GCC GAC

# FIG. 7D

Leu Glu Thr Pro Ala Asp Leu Ser Thr Ala Ile Ala Ser Arg Ser Val Ala Asp

													4.4	•		
AAG AT Lys II	C CTG e Leu	GAC Asp	TTG Leu	TAC Tyr	AAG Lys	CAT	1971 CCT Pro	GAC Asp	AAC Asn	ATC Ile	GAT Asp	GTC Val	TGG Trp	CTG Leu	GGA	998 GGC Gly
TTA GC Leu Al						AGG									TGT	
ATT GG						CTG		GAC							GAG	
AGC CA Ser Hi						CAG									CTG	
CGG GT Arg Va						GGC									TTC	
GTC GG Val Gl						TTT									ATG	
CTG GA Leu GI						TTT									CCA	
AGC GT Ser Va						GTG									GTG	
GTG TA Val Ty						TAT									ACT	
ACC CA Thr Gl	G GAA n Glu	GGA Gly	TGG Trp	GAT Asp	TTC Phe	CAG	2457 CCT Pro	CCC Pro	CTC Leu	TGC Cys	AAA Lys	GAT Asp	GTG Val	AAC Asn	GAG	484 TGT Cys

# FIG. 7E

2511 2538 GCA GAC GGT GCC CAC CCC CCC TGC CAC GCC TGT GCG AGG TGC AGA AAC ACC AAA Ala Asp Gly Ala His Pro Pro Cys His Ala Ser Ala Arg Cys Arg Asn Thr Lys 2565 2592 GGC GGC TTC CAG TGT CTC TGC GCG GAC CCC TAC GAG TTA GGA GAC GAT GGG AGA Gly Gly Phe Gin Cys Leu Cys Ala Asp Pro Tyr Glu Leu Gly Asp Asp Gly Arg 2619 2646 ACC TGC GTA GAC TCC GGG AGG CTC CCT CGG GTG ACT TGG ATC TCC ATG TCG CTG Thr Cys Val Asp Ser Gly Arg Leu Pro Arg Val Thr Trp Ile Ser Met Ser Leu 2673 2700 GCT GCT CTG CTG ATC GGA GGC TTC GCA GGT CTC ACC TCG ACG GTG ATT TGC AGG Ala Ala Leu Leu Ile Gly Gly Phe Ala Gly Leu Thr Ser Thr Val 11e Cys Arg 2727 TGG ACA CGC ACT GGC ACT AAA TCC ACA CTG CCC ATC TCG GAG ACA GGC GGA GGA Trp Thr Arg Thr Gly Thr Lys Ser Thr Leu Pro Ile Ser Glu Thr Gly Gly Gly 2781 **2808** ACT CCC GAG CTG AGA TGC GGA AAG CAC CAG GCC GTA GGG ACC TCA CCG CAG CGG Thr Pro Glu Leu Arg Cys Gly Lys His Gln Ala Val Gly Thr Ser Pro Gln Arg 2835 5865 GCC GCA GCT CAG GAC TCG GAG CAG GAG AGT GCT GGG ATG GAA GGC CGG GAT ACT Ala Ala Ala Gln Asp Ser Glu Gln Glu Ser Ala Gly Met Glu Gly Arg Asp Thr 2916 2889 CAC AGG CTG CCG AGA GCC CTC TGA GGG CAA AGT GGC AGG ACA CTG CAG AAC AGC His Arg Leu Pro Arg Ala Leu ^^^ 2970 2943 TTC ATG TTC CCA AAA TCA CCG TAC GAC TCT TTT CCA AAC ACA GGC AAA TCG GAA 3024 2997 ATC AGC AGG ACG ACT GTT TTC CCA ACA CGG GTA AAT CTA GTA CCA TGT CGT AGT 3051 TAC TOT CAG GCA TGG ATG AAT AAA TGT TAT AGC TGC AAA AAA AAA AAA

## FIG.7F

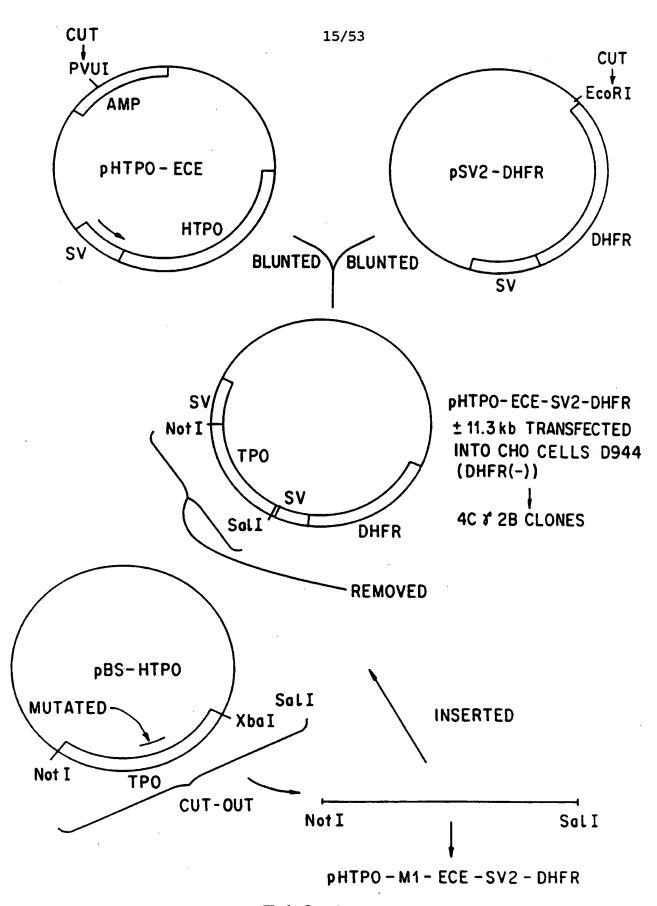
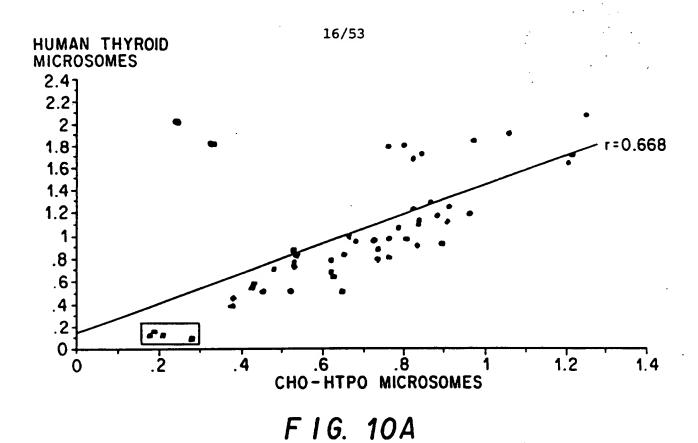
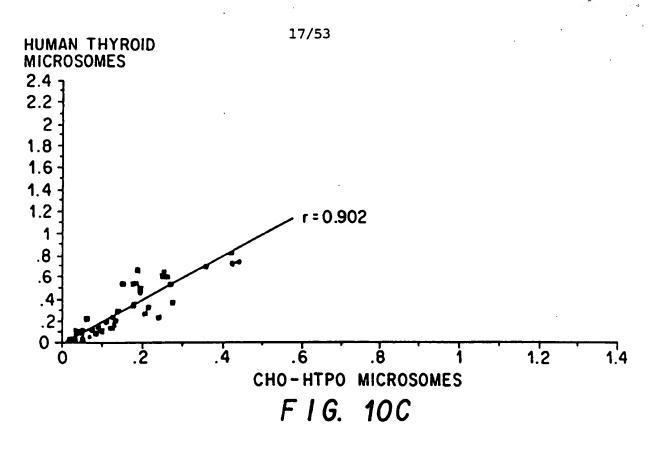
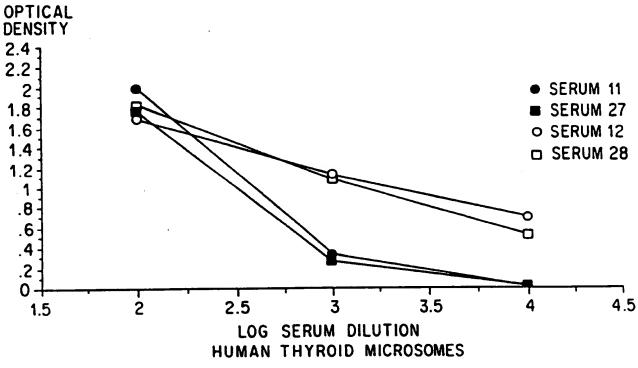


FIG. 9 SUBSTITUTE SHEET

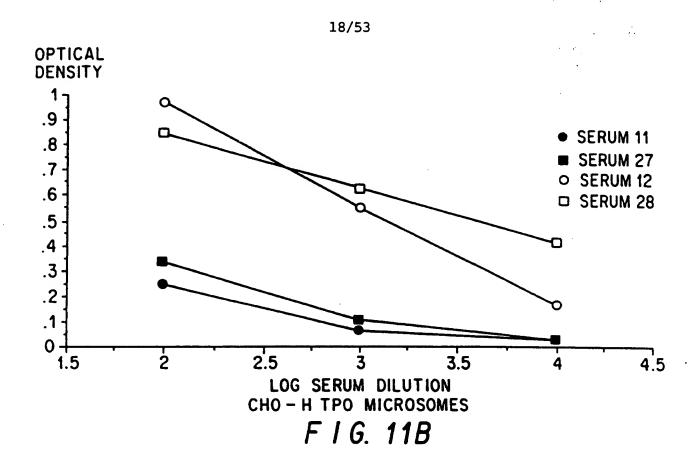


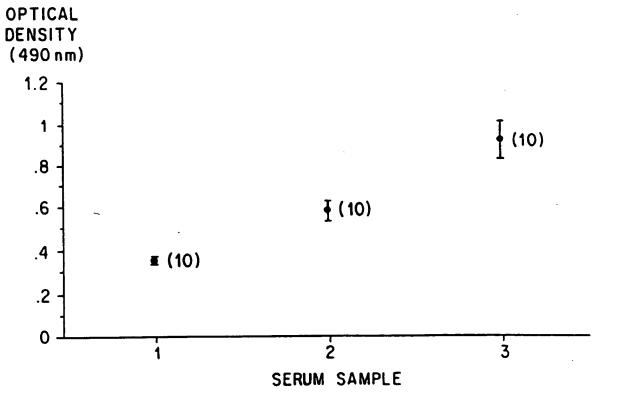
**HUMAN THYROID MICROSOMES** 2.4 2.2 2 1.8 r=0.906 1.6 1.4 1.2 1 8. .6 .4 .2 0 .4 1.2 1.4 .ż 8. .**6** CHO-HTPO MICROSOMES F I G. 10B





F | G. 11A SUBSTITUTE SHEET





**SUBSTITUTE SHEET** 

F 1 G. 12

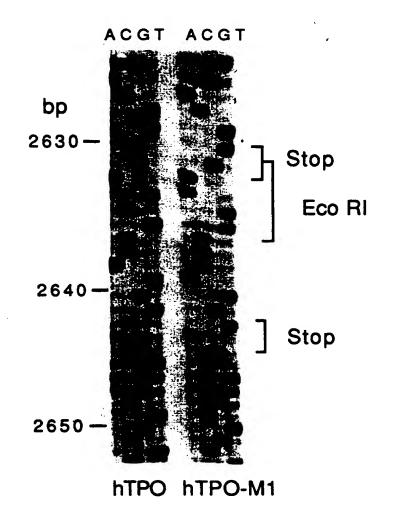
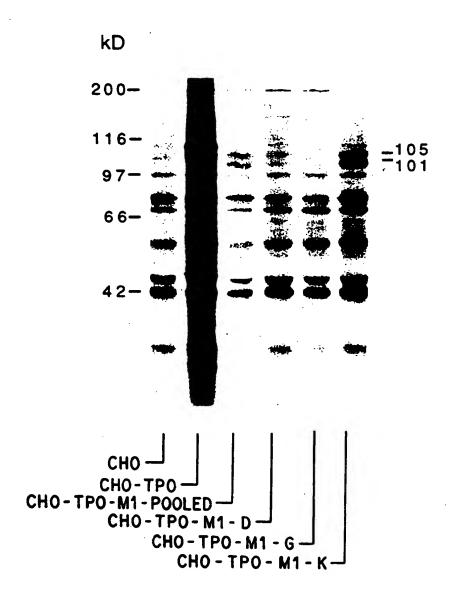
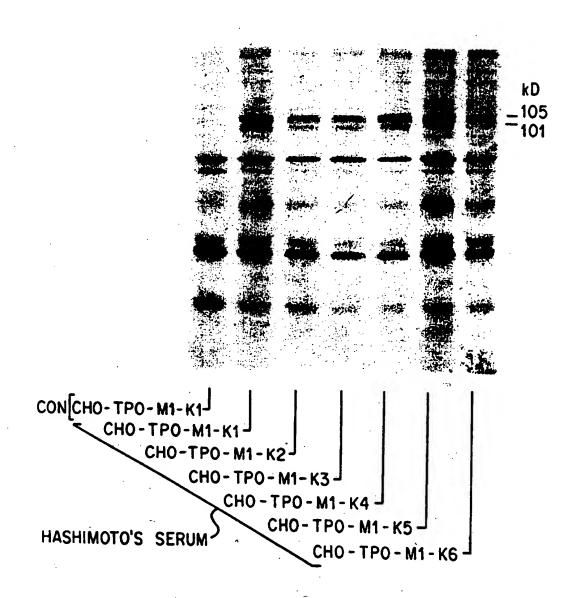


FIG. 13



F I G. 14A



F I G. 14B

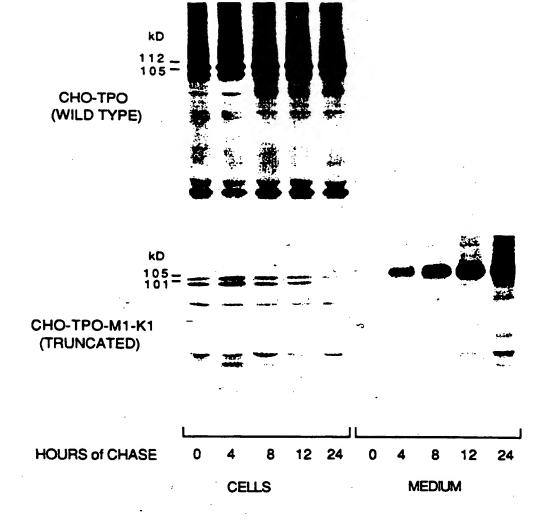
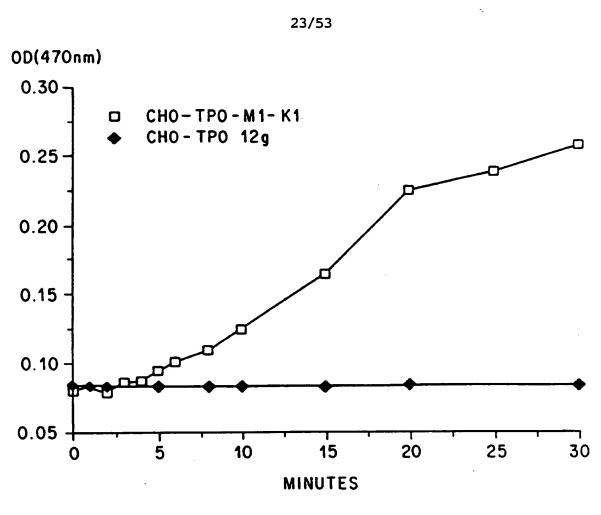
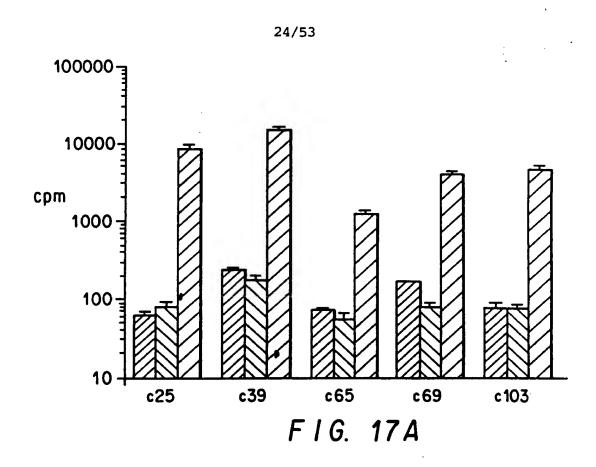
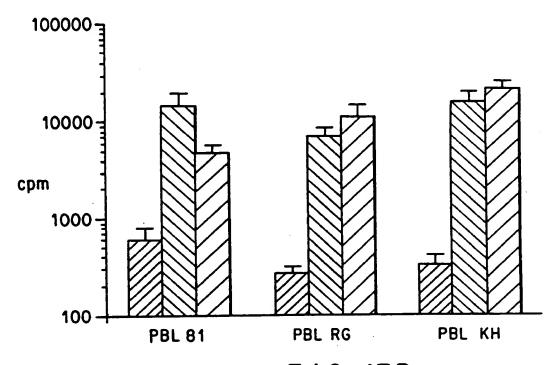


FIG. 15

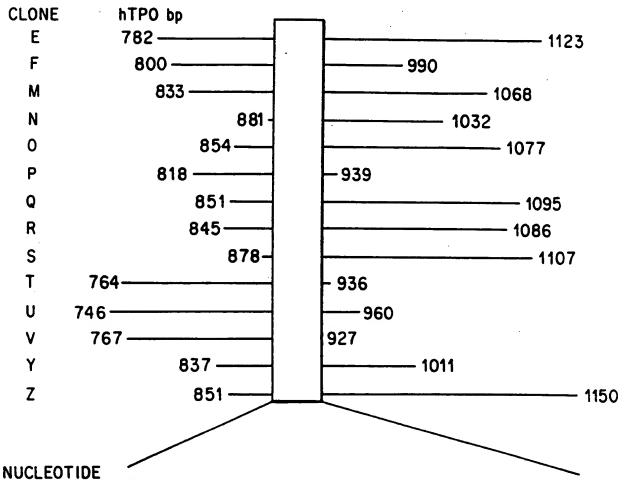


F I G. 16



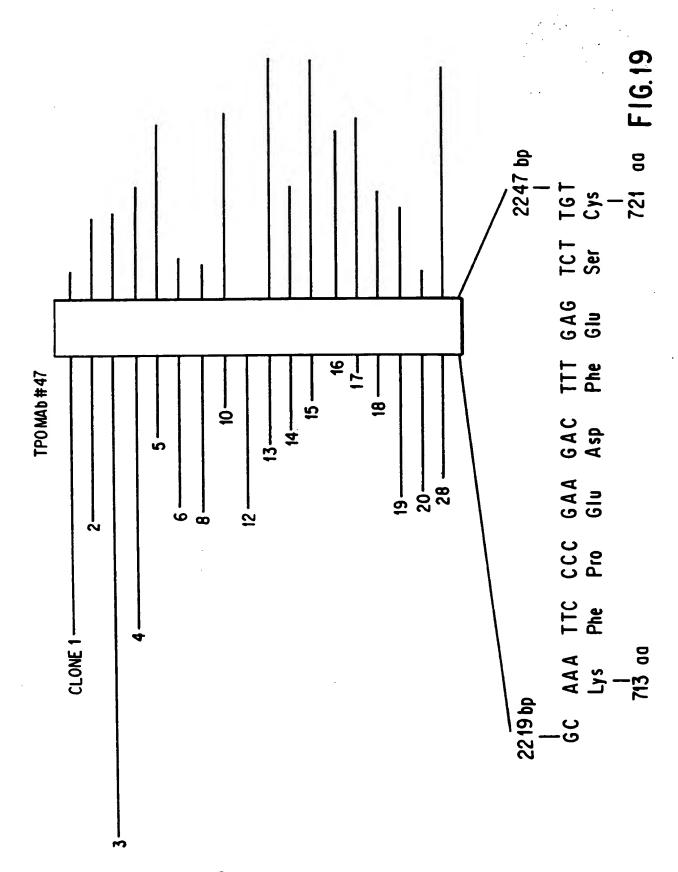


F I G. 17B SUBSTITUTE SHEET

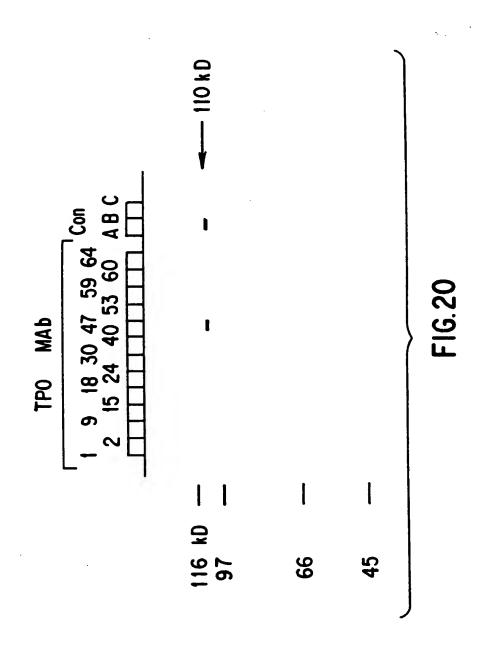


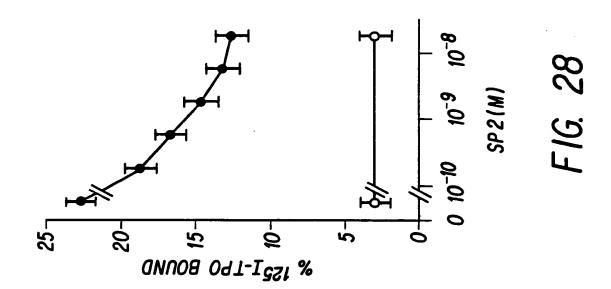
883 AAC CCA TGT TTT CCC ATA CAA CTC CCG GAG GAG GCC CGG CCG GCC 927
AMINO ACID
266 Asn Pro Cys Phe Pro Ile Gln Leu Pro Glu Glu Ala Arg Pro Ala 281

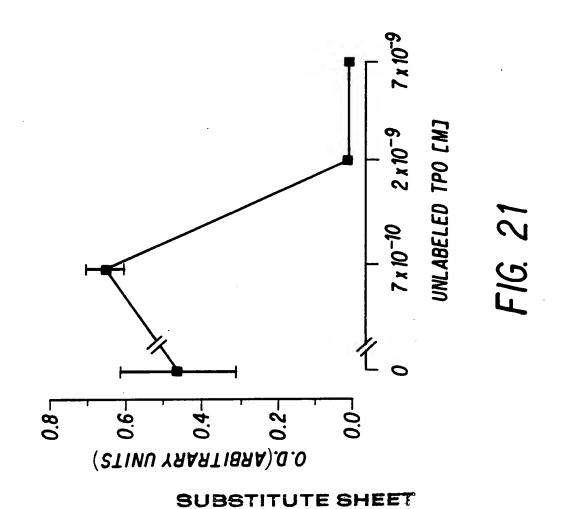
FIG. 18



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2	9	/	5	3

	. U I	120	* [H]		
VH SEGMENT (VHI)	OVKLLLESGANET VKKPGARGET VKV CAGGTGAAACTGCTCGAGTCTGAGGTGAAGAAGCCTGGGGCCTCAGTGAAGGTC CAGG-GCC	CDR1  ********  GCKARGGCTTCTGGATACACCTTCACCGGCCACTATATGCACTGGACAGGCC	CDR2  ***********************************	**************************************	MELLSGLRAGATTTGACGACACGGCCGTGTATTACTGTGCGACA
5	892 1-1	88 1.1	SUBSTIT	OTE SHE	EŢ Al

Carper F

CDR3

D SEGMENT

PCT/US92/07381

(CONT. 22 FIG.

SUBSTITUTE SHEET

SEGMENT

B

SP2

SP2 JH3

0 D H V	GAGCICGIGAIGACCCAGICICCAICTIGCTGCAICTGGAGAGAGACAGTCAGC CACA	I T C R A S E N I S R Y S N W Y Q Q Q P ATCACTTGCCGGGCAAGTGAAATATTAGCAGGTATTCAAATTGGTATCAGCAACCA TAACTAA	CDR2  ***********************************	RFSGRGGATCTGGGACACATTTCACTCTCACCATCAACAGTCTGCAACCT	•
20 时	7 4 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	* W T	CDR2  ***********************************	CATCA	
	7	**************************************	* * * * * * * * * * * * * * * * * * *	THFTL TCACTCTCAC	CDR3
L S		* * * * * * * * * * * * * * * * * * *	CDR2	FCACT	R3
I S	CICCATCTTCCCTGTCTCTCTCTCTCTCTCTCTCTCTCTCTC	**** B B B B B C B C B C B C B C B C B C B C	* * * * * * * * * * * * * * * * * * *	H I CATTI G	CDR3
80 5	R1	* * * * * * * * * * * * * * * * * * *	2017 1014 1017	GACA(	4
A C	CDR1	* * * * * COAGA	r I	rcrad	
۵ <u>.</u>		******** R A B CGGGCAAGTG	K L MAACTCO	TGGA	
		* # D	A E I	SCAG	
L V M		ν Ευ Ευ Ευ Ευ Ευ Ευ Ευ Ευ Ευ Ευ Ευ Ευ Ευ	AAGCCCT	S A GT G	
1 6	CACA	ATCACT	GK	R 4	

CDR3	G D F A T Y Y C Q Q T Y S S P F GGAGATTTTGCAACTTACTGTCAACAGACTTACAGTTCCCCGGTTC -AACC-T
	SP2 KLVJ

J SEGMENT (JK2

G Q G T K L E I K R T T GGCCAGGGACCAAGCTGAAGATCAAACGAACT

50 50 50 50	100	100 100 100	100	150 150 150 150	150
. CA		. G	CA-AGTCACC ATCA-TTGCC GGGCAAGT-A GATT-G- ATATT-AA *****************************		ATTGGTATC- GCAA-CCA GGG-AAGCCC CT-A-CT-CT-T-T-T-T-T-T-T-T-T-T-T-T-T-T-T
KLVJ VK-SP2 VK-SP4 VK-SP5 Consensus	KLVJ	VK-SP2 VK-SP4 VK-SP5	Consensus	S KIVJ KLVJ VK-SP2 VK-SP4 VK-SP5	COURGED COURSE

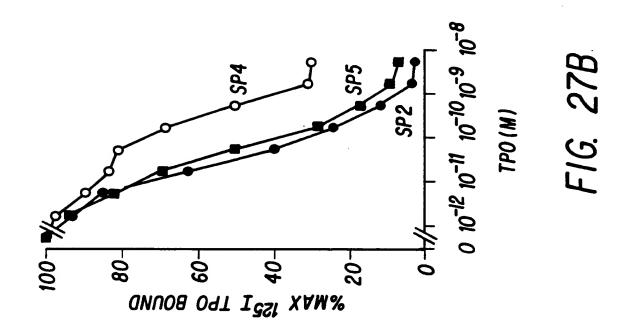
PCT/	US92.	/07381

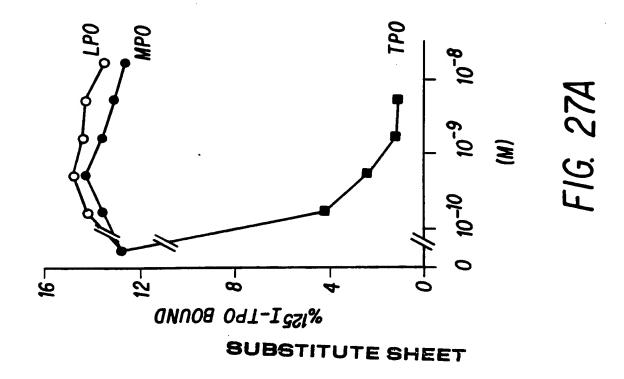
KLVJ VK-SP2 VK-SP4 VK-SP5	G		HHUH	<b>4 4 0 0</b>	700 700 700 700
Consensus	-CATCCA-TT T-CAAAGTGG	GGTCCCATCA	AGGTTCAG-G	GCAGTGG-TC	200
	**************************************				
KLVJ VK-8P2 VK-8P4 VK-8P5	0000	0 4 0 0		A B G	250 250 250 250
Consensus	TTCAC	CCATCA-CAG	TCTGCAACCT	G-Agattttg	250
KLVJ VK-SP2 VK-SP4 VK-SP5		0 0 0 0	ACCT CGTT		8 8 8 8 7 7 7 7 8 7 7 7 7
Consensus	CAACTTACTA CTGTCAACAG	CAG A-TTACAGT- ************************************	~******		<b>78</b> <b>88</b> <b>7</b>
mel	FIG.	24 (C	(CONT.)	·	

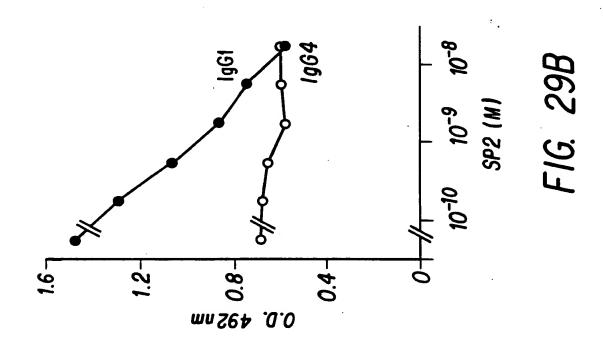
KLVJ VK-SP2 VK-SP4 VK-SP5	DI	S.S. NQTVG TR	SA E. T	50 50 50 50
Consensus	Elvmtqspss lsasvgdrvt	ITCRASQ.IYLNWYQQKP ******* CDRJ	GKAPKLLIY.	20
KLVJ VR-SP2 VR-SP4 VR-SP5	SO	D		9 6 6
Consensus	ASTLQSGVPS RFSGSGSGTD ***** CDR2	FTLTISSLQP EDFATYYC(	10 SYSTPP ****** CDR3	96
TUTE SHI	FIG	.g. 25		

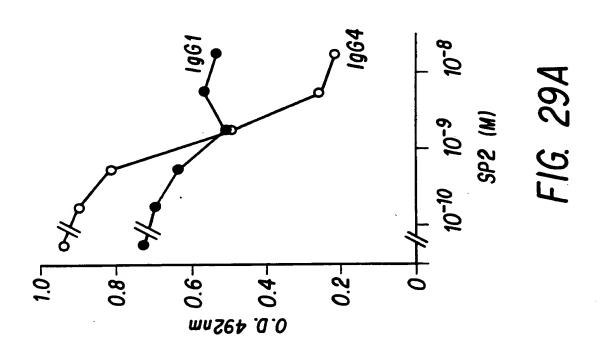
	J-KV312 JK-SP2 JK-SP4/5	E E C				<i>es es</i> e
8	Consensus		T-GGCC A-GGGACCAA	G-TGGA-ATC AAACGAACT	AAACGAACT	n en
UBST						
T	J-KV312	•	•	13		
UT	JK-SP2	•	•	13		
E :	JK-SP4/5		•	13		
SHE	Consensus	TFGQGTKLEI	KRT	13	·	
ET			ഥ	FIG. 26		

Compet

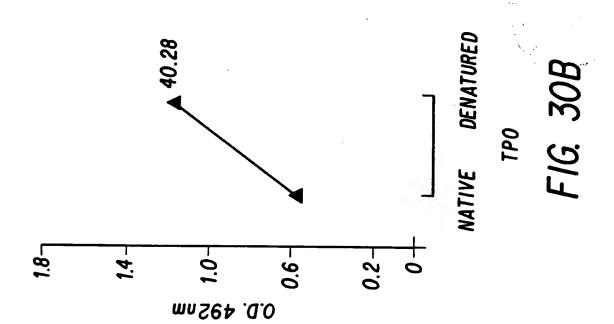


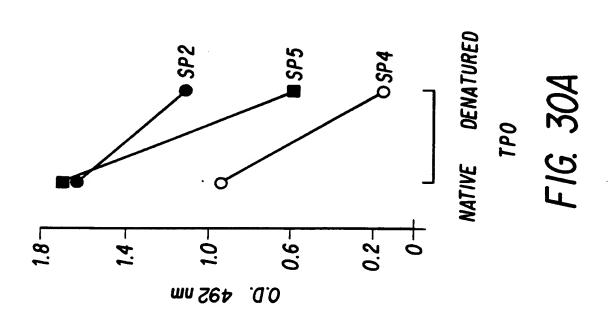






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Consensus	CAGGTG-A-C TG-TAGTC TO	TGGGGCTGAG	-TGAA-AA-C CTGGGGCCTC	CTGGGGCCTC	50
hv1L1 SP4.6	C. G G. GC		GCG.		50
Consensus hv1L1	AGTGA-GGTC TCCTGCAAGG C	TTCTGGA-A	CTTCTGGA-A CACCTTCA-C	CDR1 G-CTAC-AT-	00
SP4.6		E4	N.	ACG	00
Consensus	T-CACTGGGT GCGACAGGCC CCTGGACAAG GGCTTGAGTG G-TGGGATGG	CCTGGACAAG	GGCTTGAGTG	G-TGGGATGG	150
<b>6</b> hv1L1 <b>8</b> SP4.6	D. H.			. B	150
() TITUTE	FIG.	3. 31A	-		
mel				••	

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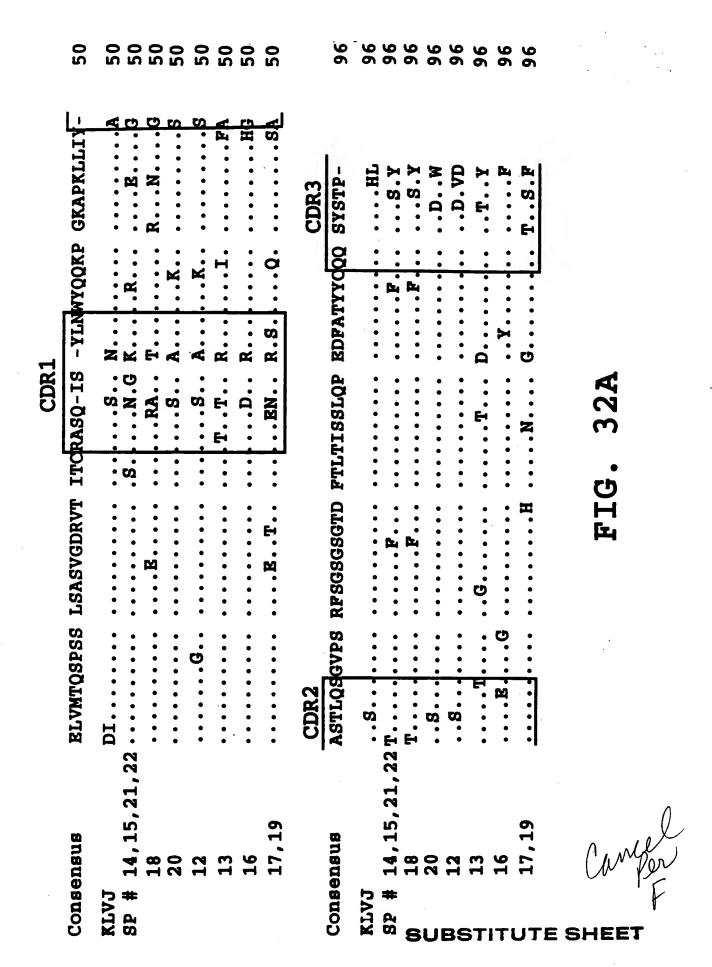
	CDR2	
Consensus	ATCAACCCTA A-A-TG-TGG CAC-Aing -C-Chears Treasggdag	200
3P4.6	G.A.C. G.BAC. G.A. T.C.	200
Consensus	GGTCACCATG ACCAG-GACA CG-CCATCAG C-CAGCCTA- ATGGAGCTGA	250
hv1L1 SP4.6		250
Consensus hv1L1 SP4.6	-CAG-CTGA- ATC-GACGAC ACGGCCGT-T A-TACTGTGC G-GA GGGTAA	2 2 2 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4
Cury	FIG. 31A (CONT.)	

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·E				CDR1		
Consensus	QV-LSGAE	PGASV-V	PGASV-V SCKASG-TFYHWVRQA PGQGLEW-GW	-YHWVRQA	PGOGLEW-GW	50
m bv1L1		VNKK.	VNKKDT G.YM	G. YM.	¥	50
- SP4.6	K.LE	LKNR.	YN D.HV	D. HV	Λ	20
	CDR2				·	
Consensus	INPGT-Y	-QKFQGRVTM	-QKFQGRVTM TRDT-IS-AY MELL-SDD TAVYYCAR	MELL-SDD	TAVYYCAR	86
hv1L1	NSGN.	<b>A</b>	AS.TSR.R	SR.R	•	98
SP4.6	KNAR.	<b>S</b>	STS.K	TS.K	•	98

GGG GTA GGA GTT GGT ACG TGG GGC CTT Ö O





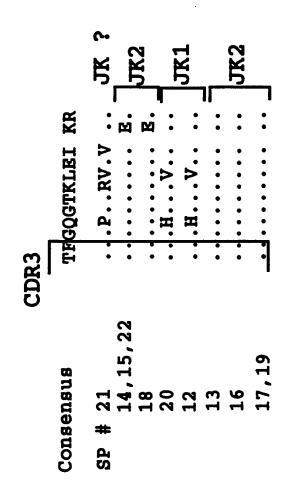


FIG. 32B

SUBSTITUTE SHEET

Carrel